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April 30, 2004

CERTIFICATE OF MAILING
37 C.F.R 1.8

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April 30, 2004 — Date
Gina N. Shishima

MS DD
Commissioner for Patents
P.O. Box 1450
Alexandria, Virginia 22313-1450

RE: *U.S. Patent Application No. 10/751,606 entitled "OPTIMIZATION OF CANCER TREATMENT WITH IRINOTECAN" – Mark J. Ratain et al.*
Our reference: ARCD:389US
Client reference: UCHI 1014

Sir:

Enclosed for filing in the above-referenced patent application is an Information Disclosure Statement, Form PTO-1449, and references A1-A8, B1-B5 and C1-C217.

No fees are believed to be due in connection with the filing of this Information Disclosure Statement, however, should any fees under 37 C.F.R. §§ 1.16 to 1.21 be deemed necessary for any reason relating to the enclosed materials, the Commissioner is authorized to deduct the appropriate fees from Fulbright & Jaworski Deposit Account No.: 50-1212/ARCD:389US.

Please date stamp and return the enclosed postcard evidencing receipt of these materials.

Respectfully submitted,

Gina N. Shishima
Reg. No. 45,104

GNS/kmv
Encl.: as noted

PATENT

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of:
Mark J. Ratain *et al.*

Serial No.: 10/751,606

Filed: January 5, 2004

For: OPTIMIZATION OF CANCER TREATMENT WITH IRINOTECAN

Group Art Unit: 1645

Examiner: Unknown

Atty. Dkt. No.: ARCD:389US

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April 30, 2004
Date

Gina N. Shishima

INFORMATION DISCLOSURE STATEMENT

MS DD
Commissioner for Patents
P.O. Box 1450
Alexandria, Virginia 22313-1450

Sir:

In compliance with the duty of disclosure under 37 C.F.R. § 1.56, it is respectfully requested that this Information Disclosure Statement be entered and the documents listed on attached Form PTO-1449 be considered by the Examiner and made of record. Copies of the listed documents required by 37 C.F.R. § 1.98(a)(2) are enclosed for the convenience of the Examiner.

In accordance with 37 C.F.R §§ 1.97(g), (h), this Information Disclosure Statement is not to be construed as a representation that a search has been made, and is not to be construed to be

an admission that the information cited is, or is considered to be, material to patentability as defined in 37 C.F.R. § 1.56(b).

The present Information Disclosure Statement is being filed prior to the receipt of a first Official Action reflecting an examination on the merits, and hence is believed to be timely filed in accordance with 37 C.F.R § 1.97(b). No fees are believed to be due in connection with the filing of this Information Disclosure Statement, however, should any fees under 37 C.F.R. §§ 1.16 to 1.21 be deemed necessary for any reason relating to these materials, the Commissioner is authorized to deduct the appropriate fees from Fulbright & Jaworski Deposit Account No.: 50-1212/ARCD:389US.

Applicants respectfully request that the listed documents be made of record in the present case.

Respectfully submitted,

Gina N. Shishima
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Attorney for Applicants

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Date: April 30, 2004

Form PTO-1449 (modified)	Atty. Docket No. ARCD:389US	Serial No. 10/751,606
List of Patents and Publications for Applicant's INFORMATION DISCLOSURE STATEMENT (Use several sheets if necessary)	Applicant Mark J. Ratain *et al.*	
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U.S. Patent Documents	Foreign Patent Documents	Other Art
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U.S. Patent Documents

Exam. Init.	Ref. Des.	Document Number	Date	Name	Class	Sub Class	Filing Date of App.
	A1	5,786,344	7/28/98	Ratain *et al.*	514	100	4/17/95
	A2	6,066,645	5/23/00	Hausheer *et al.*	514	283	1/6/99
	A3	6,287,834	9/11/01	Belanger *et al.*	435	193	2/08/99
	A4	6,319,678	11/20/01	Trubetskoy and Shaw	435	15	6/25/99
	A5	6,395,481	5/28/02	Di Rienzo *et al.*	435	6	1/16/99
	A6	6,407,117	6/18/02	Bouscarel *et al.*	514	283	3/23/00
	A7	6,472,157	10/29/02	De Rienzo and Ratain	435	6	2/01/02
	A8	6,479,236	11/12/02	Penny and Galvin	435	6	5/05/99

Foreign Patent Documents

Exam. Init.	Ref. Des.	Document Number	Date	Country	Class	Sub Class	Translation Yes/No
✓	B1	EP 0919244	6/2/99	Europe			Abstract
✓	B2	WO 00/06776	2/10/00	PCT			
✓	B3	WO 94/22846	10/94	PCT			
✓	B4	WO 95/08986	4/6/95	PCT			
	B5	WO 96/01127	1/18/96	PCT			

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	C2 ✓	Akiyama *et al.*, "Most drugs that reverse multidrug resistance also inhibit photoaffinity labeling of p-glycoprotein by a vinblastine analog," *Mol. Pharmacol.*, 33(2):144-147, 1988.

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	C7	Atsumi *et al.*, "Identification of the Metabolites of Irinotecan, a New Derivative of Camptothecin, in Rat Bile and its Biliary Excretion," *Xenobiotica*, 21(9):1159-1169, 1991.
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	C9	Barker *et al.*, "Determination of plasma concentrations of epirubicin and its metabolites by high-performance liquid chromatography during a 96-h infusion in cancer chemotherapy," *J Chromatogr B Biomed Appl*, 681:323-329, 1996.
	C10	Bear, "Drugs transported by-P-glycoprotein inhibit a 40pS outwardly rectifying chloride channel," *Biochem. Biophys. Res. Commun.*, 200(1):513-521, 1994.
	C11	Bell *et al.*, "Roles of peptidyl-prolyl cis-trans isomerase and calcineurin in the mechanisms of antimalarial action of cyclosporin A, FK506, and rapamycin," *Biochem. Pharmacol.*, 48(3):495-503, 1994.
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	C17	Bock *et al.*, In: Conjugation reactions in biotransformation, Elsevier, North Holland Biomedical Press, p. 357-364, 1978.
	C18	Boesch and Loor, "Extent and persistence of P-glycoprotein inhibition in multidrug-resistant P388 cells after exposure to resistance-modifying agents," *Anticancer Drugs*, 5(2):229-238, 1994.
	C19	Boesch *et al.*, "Restoration of daunomycin retention in multidrug-resistant P388 cells by submicromolar concentrations of SDZ PSC 833, a nonimmunosuppressive cyclosporin derivative," *Exp. Cell. Res.*, 196(1):26-32, 1991.
	C20	Boiteux-Antoine *et al.*, "Comparative induction of drug-metabolizing enzymes by hypolipidaemic compounds," *Gen-Pharmacol*, 20(4):407-412, 1989.
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	C22	Bosma *et al.*, "The genetic basis of the reduced expression of bilirubin UDP-Glucuronosyltransferase 1 in Gilbert's Syndrome," *N. Eng. J. Med.*, 333:1171-1175, 1995.
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	C25	Burger *et al.*, "Pharmacokinetic interaction between rifampin and zidovudine," *Antimicrobial Agents and Chemotherapy*, 37(7):1426-1431, 1993.
	C26	Campain *et al.*, "Characterization of an unusual mutant of human melanoma cells resistant to anticancer drugs that inhibit topoisomerase II," *J. Cell Physiol.*, 155(2):414-425, 1993.
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	C29	Cascorbi et al., "Frequency of single nucleotide polymorphisms in the p-glycoprotein drug transporter MDR1 gene in white subjects," *Clinc. Pharmacol Ther.*, 69:169-174, 2001.
	C30	Charuk *et al.*, "Interaction of Rat Kidney P-Glycoprotein with a Urinary Component and Various Drugs Including Cyclosporin A," *Am. J. Physiol.*, 266:F66-F75, 1994.
	C31	Chen *et al.*, "Fluorescence polarization in homogeneous nucleic acid analysis," *Genome Res.*, 9:492-498, 1999.
	C32	Chen *et al.*, "Calcium phosphate-mediated gene transfer: A highly efficient transfection system for stably transforming cells with plasmid DNA," *Biotechniques*, 6:632-638, 1988.
	C33	Cheng *et al.*, "Glucuronidation of catechol estrogens by expressed human UDP-glucuronosyltransferases (UGTs) 1A1, 1A3, and 2B7," *Toxicological Sciences*, 45:52-57, 1998.
	C34	Chien *et al.*, "In vitro evaluation of flavopiridol, a novel cell cycle inhibitor, in bladder cancer," *Cancer Chemother Pharmaco*, 44:81-87, 1999.
	C35	Chin *et al.*, "Reduced mRNA levels for multidrug-resistance genes in cAMP-dependent protein kinase mutant cell lines," *J. Cell Physiol.*, 152(1):87-94, 1992.
	C36	Clarke and Burchell, "The Uridine Diphosphate glucuronosyltransferase multigene family: function and regulation," *Handbook of experimental pharmacology*, 112:3-43, 1994.
	C37	Coffman *et al.*, "Cloning and stable expression of a cDNA encoding a rate liver UDP-Glucuronosyltransferase (UDP_Glucuronosyltransferase 1.1) that catalyzes the glucuronidation of opiods and bilirubin," *Mol. Pharmacol.*, 47:1101-1105, 1995.
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	C39	Coffman *et al.*, "The glucuronidation of opioids, other xenobiotics, and androgens by human UGT2B7Y(268) and UGT2B7H(268)," *Drug Metab Dispos*, 26:73-77, 1998.
	C40	Cordon-Cardo *et al.*, "Expression of the multidrug resistant gene product (P-glycoprotein) in human normal and tumor tissues," *J. Histochem. Cytochem.*, 38:1277-1287, 1990.
	C41	Czech *et al.*, "Antitumoral activity of flavone L86-8275," *Int J Oncol*, 6:31-66, 1995.

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	C45	De Morais *et al.*, "Biotransformation and Toxicity of Acetaminophen in Congenic RHA Rats with or without a Hereditary Deficiency in Bilirubin UDP-Glucuronosyltransferase," *Toxicology and Applied Pharmacology*, 117:81-87, 1992.
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	C47	Dhainaut *et al.*, "New Triazine Derivatives as Potent Modulators of Multidrug Resistance," *J. Med. Chem.*, 35:2481-2496, 1992.
	C48	Di Carlo *et al.*, "Flavonoids: old and new aspects of a class of natural therapeutic drugs," *Life Sci*, 65:337-353, 1999.
	C49	Di Rienzo *et al.*, "Two new alleles in the promoter of the bilirubin UDP-glucuronosyl transferase 1 (UGT1A1) gene", *American Society for Clinical Pharmacology and Therapeutics*, Ninety Ninth Annual Meeting, New Orleans, Abstract OII-B-3, page 207, 1998.
	C50	Diasio et al., "Clinical pharmacology of 5-fluorouracil," *Clin Pharmacokinet,* 16:215-237, 1989.
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	C52	Doige *et al.*, "ATPase activity of partially purified P-glycoprotein from multidrug-resistant chinese hamster ovary cells," *Biochim. Biophys. Acta.*, 1109(2):149-160, 1992.
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	C54	Egner *et al.*, "Regulation of Phase 2 Enzyme Induction by Oltipraz and other Dithiolethiones," *Carcinogenesis*, 15(2):177-181, 1994.

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	C56	Evans and Relling, "Automated high-performance liquid chromatographic assay for the determinationof 7-ethoxycoumarin and umbelliferone," *J. Chromatogr.*, 578:141-145, 1992.
	C57	Ford *et al.*, "Cellular and biochemical characterization of thioxanthenes for reversal of multidrug resistance in human and murine cell lines," *Cancer Res.*, 50(6):1748-1756, 1990.
	C58	Fournel *et al.*, "Structure-dependent induction of bilirubin glucuronidation and lauric acid 12-hydroxylation by arylcarboxylic acids chemically related to clofibrate," *Biochimica at Biophysica Acta*, 842:202-213, 1985.
	C59	Foxwell *et al.*, "Identification of the multidrug resistance-related P-glycoprotein as a cyclosporine binding protein," *Mol. Pharmacol.*, 36:543-546, 1989.
	C60	Friche *et. al.*, "In vitro circumvention of anthracycline-resistance in ehrlich ascites tumour by anthracycline analogues" *Biochem. Pharmacol.*, 39:1721-1726, 1990.
	C61	GenBank Accession Number AF297093.
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	C63	Gestl *et al.*, "Expression of UGT2B7, a UDP-glucuronosyltransferase implicated in the metabolism of 4-hydroxyestrone and all-trans retinoic acid, in normal human breast parenchyma and in invasive and in Situ breast cancers," *American Journal of Pathology*, 160(4):1467-1479, 2002.
	C64	Gram *et al.*, "Clinical relevance of gentic polymorphisms in drug oxidation," *Clinical Relevance of Genetic Polymorphisms in Drug Oxidation*, 1992.
	C65	Green *et al.*, "Expressed human UGT1.4 protein catalyzes the formation of quaternary ammonium-linked glucuronides," *Drug Metab. Dispos.*, 23:299-302, 1995.
	C66	Gruol *et al.*, "Reversal of multidrug resistance by RU 486[1]" *Cancer Res.*, 54(12):3088-3091, 1994.
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	C74	Gutmann *et al.*, "Modulation of multidrug resistance protein expression in porcine brain capillary endothelial cells in vitro," *Drug Metab Dispos.* 27:937-941, 1999.
	C75	Hait *et al.*, "Terferadine (seldane®): a new drug for restoring sensitivity to multidrug resistant cancer cells" *Biochem. Pharmacol.*, 45(2):401-406, 1993.
	C76	Hamada *et al.*, "Mouse-human chimeric antibody against the multidrug transporter P-glycoprotein" *Cancer Res.*, 50(11):3167-3171, 1990.
	C77	Harding *et al.*, "Cloning and substrate specificity of a human phenol UDP-glucuronosyltransferase expressed in COS-7 cells," *PNAS, USA*, 85:8381-8385, 1988.
	C78	Hecht *et al.*, "4-(Methylnitrosamino)-1-(3-pyridyl)-1-butanol (NNAL) and its glucuronide, metabolites of a tobacco-specific lung carcinogen, in the urine of black and white smokers," *Proceedings of the American Association for Cancer Research*, 35:1702, 1994.
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	C80	Hjelle, "Hepatic UDP-Glucuronic Acid Regulation during Acetaminophen Biotransformation in Rats," *The Journal of Pharmacology and Experimental Therapeutics*, 237(3):750-756, 1986.
	C81	Hoffmeyer *et al.*, "Functional polymorphisms of the human multidrug-resistance gene: multiple sequence variations an dcorrelation of one allele with p-glycoprotein expression and activity in vivo," *PNAS*, 28:97(7):3473-3478, 2000.

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Form PTO-1449 (modified)	Atty. Docket No. ARCD:389US	Serial No. 10/751,606
List of Patents and Publications for Applicant's INFORMATION DISCLOSURE STATEMENT	Applicant Mark J. Ratain *et al.*	
(Use several sheets if necessary)	Filing Date: January 5, 2004	Group: 1645

U.S. Patent Documents	Foreign Patent Documents	Other Art
See Page 1	*See Page 1*	*See Page 1*

Other Art (Including Author, Title, Date Pertinent Pages, Etc.)

Exam. Init.	Ref. Des.	Citation
	C196	Stinson *et al.*, "Determination of flavopiridol (L86 8275; NSC 649890) in human plasma by reversed-phase liquid chromatography with electrochemical detection," *Cancer Chemother. Pharmacol.*, 42(4):261-265, 1998.
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